### **Amendments to the Claims:**

Please enter the following claim amendments:

1. (Currently Amended) A compound of formula (I)

$$Ar^{1} - CHCH_{2}NHCR^{4}R^{5}(CH_{2})_{m} - O - (CH_{2})_{n}$$

$$OH$$

$$(I)$$

or a salt, solvate, or physiologically functional derivative thereof, wherein:

m is an integer of from 2 to 8; n is an integer of from 3 to 11<del>, preferably from 3 to 7</del>; with the proviso that m + n is 5 to 19<del>, preferably from 5 to 12</del>;

R<sup>1</sup> is -XNR<sup>6</sup>C(O)NR<sup>7</sup>R<sup>8</sup>; wherein

X is selected from  $-(CH_2)_p$ - and  $C_{2-6}$ alkenylene;

 $R^6$  and  $R^8$  are independently selected from hydrogen,  $C_{1-6}$ alkyl and  $C_{3-7}$  cycloalkyl; wherein said  $C_{1-6}$ alkyl and  $C_{3-7}$  cycloalkyl moieties may optionally be substituted by  $-CO_2H$  or  $-CO_2(C_{1-4})$ alkyl;

 $R^7$  is selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{3-7}$ cycloalkyl,  $-C(O)R^9$ , phenyl, naphthyl, hetaryl, and phenyl( $C_{1-4}$ alkyl)- and  $R^7$  is optionally substituted by 1 or 2 groups independently selected from halo, hydroxy,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl) and  $-CO_2(C_{1-4}$ alkyl) and  $-CO_2(C_{1-4}$ alkyl) and  $-CO_2(C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl),  $-SO_2(C_{1-6}$ alkyl)

 $R^9$  is selected from  $C_{1-6}$ alkyl,  $C_{3-7}$ cycloalkyl,  $-CO_2H$ ,  $CO_2(C_{1-4}$ alkyl), phenyl, naphthyl, hetaryl, and phenyl( $C_{1-4}$ alkyl)- and  $R^9$  is optionally substituted by 1

or 2 groups independently selected from halo,  $C_{1-6}$ alkyl,  $C_{1-6}$ haloalkyl,  $C_{1-6}$ alkoxy, -NHC(O)( $C_{1-6}$ alkyl), -SO<sub>2</sub>( $C_{1-6}$ alkyl), -SO<sub>2</sub>(phenyl), -CO<sub>2</sub>H, and - CO<sub>2</sub>( $C_{1-4}$ alkyl);

 $R^{10}$  and  $R^{11}$  each independently represent hydrogen,  $C_{1-4}$ alkyl or  $C_{3-7}$  cycloalkyl, and

p is an integer from 0 to 6, preferably from 0 to 4;

or R<sup>1</sup> is cyclised such that R<sup>8</sup> forms a bond with the phenyl ring to which R<sup>1</sup> is attached, via the ring carbon atom adjacent to R<sup>1</sup>, so as to form a moiety of the formula:

 $R^2$  is selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkoxy, phenyl, halo, and  $C_{1-6}$ haloalkyl;

 $R^3$  is selected from hydrogen, hydroxy,  $C_{1-6}$ alkyl, halo,  $C_{1-6}$ alkoxy, phenyl,  $C_{1-6}$ haloalkyl, and  $-SO_2NR^{12}R^{13}$ ;

wherein  $R^{12}$  and  $R^{13}$  are independently selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{3-6}$ cycloalkyl, phenyl, and phenyl ( $C_{1-4}$ alkyl), or  $R^{12}$  and  $R^{13}$ , together with the nitrogen to which they are bonded, form a 5-, 6-, or 7- membered nitrogen containing ring;

and R<sup>12</sup> and R<sup>13</sup> are each optionally substituted by one or two groups selected from halo, C<sub>1-6</sub>alkyl, and C<sub>1-6</sub>haloalkyl;

R<sup>4</sup> and R<sup>5</sup> are independently selected from hydrogen and C<sub>1-4</sub>alkyl with the proviso that the total number of carbon atoms in R<sup>4</sup> and R<sup>5</sup> is not more than 4;

and Ar1 is a group selected from

$$R^{14}$$
 $R^{16}$ 
 $R^{16}$ 
 $R^{16}$ 
 $R^{17}$ 
 $R^{17}$ 
 $R^{17}$ 
 $R^{17}$ 
 $R^{17}$ 
 $R^{17}$ 
 $R^{19}$ 
 $R$ 

wherein R<sup>14</sup> represents hydrogen, halogen, -(CH<sub>2</sub>)<sub>q</sub>OR<sup>18</sup>, -NR<sup>18</sup>C(O)R<sup>19</sup>, -NR<sup>18</sup>SO<sub>2</sub>R<sup>19</sup>, -SO<sub>2</sub>NR<sup>18</sup>R<sup>19</sup>, -NR<sup>18</sup>R<sup>19</sup>, -OC(O)R<sup>20</sup> or OC(O)NR<sup>18</sup>R<sup>19</sup>, and R<sup>15</sup> represents hydrogen, halogen or C<sub>1-4</sub> alkyl;

or R<sup>14</sup> represents –NHR<sup>21</sup> and R<sup>15</sup> and –NHR<sup>21</sup> together form a 5- or 6-membered heterocyclic ring;

R<sup>16</sup> represents hydrogen, halogen, –OR<sup>18</sup> or –NR<sup>18</sup>R<sup>19</sup>;

 $R^{17}$  represents hydrogen, haloC<sub>1-4</sub> alkyl, -OR<sup>18</sup>, -NR<sup>18</sup>R<sup>19</sup>, -OC(O)R<sup>20</sup> or OC(O)NR<sup>18</sup>R<sup>19</sup>;

 $\mathsf{R}^{18}$  and  $\mathsf{R}^{19}$  each independently represents hydrogen or  $\mathsf{C}_{1\text{-}4}$  alkyl, or in the groups

–NR<sup>18</sup>R<sup>19</sup>, -SO<sub>2</sub>NR<sup>18</sup>R<sup>19</sup> and –OC(O)NR<sup>18</sup>R<sup>19</sup>, R<sup>18</sup> and R<sup>19</sup> independently represent hydrogen or C<sub>1-4</sub> alkyl or together with the nitrogen atom to which they are attached form a 5-, 6- or 7- membered nitrogen-containing ring,

 $R^{20}$  represents an aryl (eg phenyl or naphthyl) group which may be unsubstituted or substituted by one or more substituents selected from halogen,  $C_{1-4}$  alkyl,

hydroxy, C<sub>1-4</sub> alkoxy or halo C<sub>1-4</sub> alkyl; and

q is zero or an integer from 1 to 4;

provided that in the group (a) when  $R^{14}$  represents  $-(CH_2)_qOR^{18}$  and q is 1,  $R^{16}$  is not OH.

2. (Currently Amended) A compound of formula (I) as defined in claim 1 wherein  $R^6$  and  $R^8$  are independently selected from hydrogen,  $C_{1-6}$ alkyl and  $C_{3-7}$  cycloalkyl;

 $R^7$  is selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{3-7}$ cycloalkyl,  $-C(O)R^9$ , phenyl, naphthyl, hetaryl, and phenyl( $C_{1-4}$ alkyl)- and  $R^7$  is optionally substituted by 1 or 2 groups independently selected from halo, hydroxy,  $C_{1-6}$ alkyl,

 $C_{1-6}$ haloalkyl,  $C_{1-6}$  alkoxy, -NHC(O)( $C_{1-6}$ alkyl), -SO<sub>2</sub>( $C_{1-6}$ alkyl), -SO<sub>2</sub>(phenyl),

-CO<sub>2</sub>H, and -CO<sub>2</sub>(C<sub>1-4</sub>alkyl);

R14 is selected from the group consisting of halogen, -(CH<sub>2</sub>)<sub>q</sub>OR<sup>18</sup>,

 $-{\sf NR^{18}C(O)R^{19}, -NR^{18}SO_2R^{19}, -SO_2NR^{18}R^{19}, -NR^{18}R^{19}, -OC(O)R^{20},}$ 

-OC(O)NR<sup>18</sup>R<sup>19</sup>, alkyl, –NHR<sup>21</sup>, and R<sup>15</sup> and –NHR<sup>21</sup> together form a 5- or 6-membered heterocyclic ring;

R<sup>14</sup>-is as defined above except that R<sup>14</sup>-does not represent hydrogen; and all other substituents are as defined for formula (I).

or a salt, solvate or physiologically functional derivative thereof.

- 3. (Currently Amended) A compound according to claim 1 er claim 2 wherein R<sup>14</sup> represents hydrogen, halogen, -NR<sup>18</sup>C(O)R<sup>19</sup>, -NR<sup>18</sup>SO<sub>2</sub>R<sup>19</sup>, -SO<sub>2</sub>NR<sup>18</sup>R<sup>19</sup>, -NR<sup>18</sup>R<sup>19</sup>, -OC(O)R<sup>20</sup> or OC(O)NR<sup>18</sup>R<sup>19</sup>; and R<sup>16</sup> represents hydrogen, halogen, -OR<sup>18</sup> or -NR<sup>18</sup>R<sup>19</sup>.
- 4. (Currently Amended) A compound according to claim 1 or claim 2 wherein  $R^{14}$  represents hydrogen, halogen, -(CH<sub>2</sub>)<sub>q</sub>OR<sup>18</sup>, -NR<sup>18</sup>C(O)R<sup>19</sup>, -NR<sup>18</sup>SO<sub>2</sub>R<sup>19</sup>, -SO<sub>2</sub>NR<sup>18</sup>R<sup>19</sup>, -NR<sup>18</sup>R<sup>19</sup>, -OC(O)R<sup>20</sup> or OC(O)NR<sup>18</sup>R<sup>19</sup>; and R<sup>16</sup> represents hydrogen, halogen, or –NR<sup>18</sup>R<sup>19</sup>.
- 5. (Currently Amended) A compound of formula (I) according to <u>claim 1</u> any of claims 1 to 4 wherein R¹ represents –(CH₂)₀NHC(O)NHR³.
- 6. (Currently Amended) A compound according to <u>claim 1</u> any of claims 1 to 5 wherein p is 0, 1 or 2.
- 7. (Currently Amended) A compound of formula (I) which is selected from:

N-[3-(4-{[6-({(2R)-2-[3-(Formylamino)-4-hydroxyphenyl]-2-hydroxyethyl}amino)hexyl]oxy}butyl)phenyl]urea;
N-[3-(4-{[6-({(2R)-2-[3-(Formylamino)-4-hydroxyphenyl]-2-hydroxyethyl}amino)hexyl]oxy}butyl)phenyl]-N'-phenylurea;
N-[3-(4-{[6-({(2R)-2-[3-(Formylamino)-4-hydroxyphenyl]-2-hydroxyethyl}amino)hexyl]oxy}butyl)phenyl]-N'-pyridin-3-ylurea;
N-[3-(4-{[6-({2-hydroxy-2-[5-hydroxy-6-(hydroxymethyl)pyridin-2-

yl]ethyl}amino)hexyl]oxy}butyl)-5-methylphenyl]urea.

and salts, solvates, and physiologically functional derivatives thereof.

8. (Currently Amended) A method for the prophylaxis or treatment of a clinical condition in a mammal, such as a human, for which a selective  $\beta_2$ -adrenoreceptor agonist is indicated, which comprises <u>administrating</u> administration of a therapeutically effective amount of a compound of formula (I), (Ia) or (Ib) according to <u>claim 1</u> any of claims 1 to 7, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof.

#### 9. (Canceled)

- 10. (Currently Amended) A pharmaceutical formulation comprising a compound of formula (I) according to <u>claim 1</u> any of claims 1 to 7 or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof, and a pharmaceutically acceptable carrier or excipient, and optionally one or more other therapeutic ingredients.
- 11. (Currently Amended) A combination comprising a compound of formula (I) according to <u>claim 1</u> any of claims 1 to 7 or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof, and one or more other therapeutic ingredients.
- 12. (Original) A combination according to claim 11 wherein the other therapeutic ingredient is a corticosteroid, an anticholinergic or a PDE4 inhibitor.

#### 13. (Canceled)

14. (Currently Amended) A process for the preparation of a compound of formula (I) according to <u>claim 1</u> any of claims 1 to 7, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

# (a) deprotection of deprotecting a protected intermediate, for example of formula (II):

$$Ar^{1a} - CHCH_2NP^2CR^4R^5(CH_2)_m - O - (CH_2)_n$$

$$OP^1$$

$$R^3$$
(II)

or a salt or solvate thereof, wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m, and n are as defined for the compound of formula (I), Ar<sup>1a</sup> represents an optionally protected form of Ar<sup>1</sup>; and P<sup>1</sup> and P<sup>2</sup> are each independently either hydrogen or a protecting group, provided that the compound of formula (II) contains at least one protecting group.

## (b) alkylation of an amine of formula (IX)

wherein Ar<sup>1a</sup> is an optionally protected form of Ar<sup>1</sup> and P<sup>2</sup> is either hydrogen or a protecting group,

#### with a compound of formula (X):

$$L^{1}CR^{4}R^{5}(CH_{2})_{III} - O - (CH_{2})_{II}$$
 (X)

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m, and n are as defined for the compound of formula (I) or (Ia) and L<sup>1</sup> is a leaving group;

# (c) reduction of a compound of formula (XII):

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m and n are as defined for formula (I), Ar<sup>1a</sup> is an optionally protected form of Ar<sup>1</sup>, and P<sup>1</sup> and P<sup>2</sup> are each independently hydrogen or a protecting group as defined above;

## (d) reacting a compound of formula (XVI):

wherein Ar<sup>1a</sup> is an optionally protected form of Ar<sup>1</sup>, and P<sup>1</sup> is hydrogen or a protecting group and L<sup>4</sup> is a leaving group as defined above for groups L-L<sup>3</sup> or a compound of formula (XVII):

wherein Ar13 is as hereinbefore defined with an amine of formula (XVIII):

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, P<sup>2</sup>, m and n are as defined for formula (II); or

(e) removal of a chiral auxiliary from a compound of formula (IIa):

wherein R<sup>1</sup>—R<sup>5</sup>, m and n are as defined for formula (I), Ar<sup>1a</sup> and P<sup>1</sup> are as defined for formula (II) each independently represent hydrogen or a protecting group and R<sup>28</sup> represents a chiral auxiliary.

optionally followed by one or more of the following steps in any order selected from the group consisting of:

- (i) optional removal of removing any protecting groups;
- (ii) optional separation of separating an enantiomer from a mixture of enantiomers; and
- (iii) optional conversion of converting the product to a corresponding salt, solvate, or physiologically functional derivative thereof.
- 15. (New) A compound of formula (I) according to claim 1, wherein n ranges from 3 to 7.

- 16. (New) A compound of formula (I) according to claim 1, wherein m + n ranges from 5 to 12.
- 17. (New) A compound of formula (I) according to claim 1, wherein p ranges from 0 to 6.
- 18. (New) A compound of formula (I) according to claim 1, wherein R<sup>20</sup> represents a phenyl group.
- 19. (New) A compound of formula (I) according to claim 1, wherein R<sup>20</sup> is a naphthyl group.
- 20. (New) A method according to claim 8, wherein the mammal is a human.
- 21. (New) A method according to claim 8, wherein the clinical condition is asthma.
- 22. (New) A method according to claim 8, wherein the clinical condition is COPD.
- 23. (New) A process for the preparation of a compound of formula (I) according to claim 1, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

alkylating an amine of formula (IX)

wherein  $Ar^{1a}$  is an optionally protected form of  $Ar^1$  and  $P^2$  is either hydrogen or a protecting group,

with a compound of formula (X):

$$L^{1}CR^{4}R^{5}(CH_{2})_{m} -O -(CH_{2})_{n}$$
 $R^{2}$ 
 $R^{1}$ 
 $(X)$ 

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m, and n are as defined for the compound of formula (I) and L<sup>1</sup> is a leaving group;

optionally followed by one or more of the following steps in any order selected from the group consisting of:

- (i) removing any protecting groups;
- (ii) separating an enantiomer from a mixture of enantiomers; and
- (iii) converting the product to a corresponding salt, solvate, or physiologically functional derivative thereof.
- 24. (New) A process for the preparation of a compound of formula (I) according to claim 1, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

reducing a compound of formula (XII):

$$Ar^{1a} - CHCH_2NP^2CR^4R^5(CH_2)_m - O - (CH_2)_{n-2} = R^2$$

$$| \qquad \qquad |$$

$$OP^1 \qquad \qquad |$$
(XII)

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m and n are as defined for formula (I), Ar<sup>1a</sup> is an optionally protected form of Ar<sup>1</sup>, and P<sup>1</sup> and P<sup>2</sup> are each independently hydrogen or a protecting group;

optionally followed by one or more of the following steps in any order selected from the group consisting of:

- (i) removing any protecting groups;
- (ii) separating an enantiomer from a mixture of enantiomers; and
- (iii) converting the product to a corresponding salt, solvate, or physiologically functional derivative thereof.
- 25. (New) A process for the preparation of a compound of formula (I) according to claim 1, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

reacting a compound of formula (XVI):

wherein Ar<sup>1a</sup> is an optionally protected form of Ar<sup>1</sup>, and P<sup>1</sup> is hydrogen or a protecting group, and L<sup>4</sup> is a leaving group or a compound of formula (XVII):

wherein Ar<sup>1a</sup> is as hereinbefore defined with an amine of formula (XVIII):

$$P^2HNCR^4R^5(CH_2)_m$$
 —  $O$  —  $(CH_2)_n$   $R^3$  (XVIII)

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m and n are as defined for formula (I) and P<sup>2</sup> is hydrogen or a protecting group; or

optionally followed by one or more of the following steps in any order selected from the group consisting of:

- (i) removing any protecting groups;
- (ii) separating an enantiomer from a mixture of enantiomers; and
- (iii) converting the product to a corresponding salt, solvate, or physiologically functional derivative thereof.
- 26. (New) A process for the preparation of a compound of formula (I) according to claim 1, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

removal of a chiral auxiliary from a compound of formula (IIa):

$$Ar^{1a} - CHCH_2NR^{28}CR^4R^5(CH_2)_m - O - (CH_2)_n$$

$$OP^1 \cdot (IIa)$$

wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m and n are as defined for formula (I), Ar<sup>1a</sup> and P<sup>1</sup> each independently represent hydrogen or a protecting group and R<sup>28</sup> represents a chiral auxiliary

optionally followed by one or more of the following steps in any order selected from the group consisting of:

- (i) removing any protecting groups;
- (ii) separating an enantiomer from a mixture of enantiomers; and
- (iii) converting the product to a corresponding salt, solvate, or physiologically functional derivative thereof.